

L35 ANSWER 190 OF 272 HCAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1993:225657 HCAPLUS Full-text
 DOCUMENT NUMBER: 118:225657
 TITLE: Cyclosporin immunoassay and reagents
 INVENTOR(S): Davalian, Dariush; Beresini, Maureen H.; Alexander, Svetlana; Hu, Mae W.; Ullman, Edwin F.
 PATENT ASSIGNEE(S): Syntex (U.S.A.), Inc., USA
 SOURCE: Eur. Pat. Appl., 56 pp.
 CODEN: EPXXDW
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 487289	A2	19920527	EP 1991-310632	19911119
EP 487289	A3	19940223		
EP 487289	B1	19960904		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
CA 2055813	AA	19920521	CA 1991-2055813	19911119
EP 674178	A2	19950927	EP 1995-108148	19911119
EP 674178	A3	19960710		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE				
AT 142344	E	19960915	AT 1991-310632	19911119
ES 2091301	T3	19961101	ES 1991-310632	19911119
ES 2181734	T3	20030301	ES 1995-108148	19911119
JP 04316599	A2	19921106	JP 1991-354242	19911120
JP 3433952	B2	20030804		
JP 2003201298	A2	20030718	JP 2002-366051	19911120
US 6054303	A	20000425	US 1995-401827	19950310
US 6190873	B1	20010220	US 1995-402296	19950310
US 6410696	B1	20020625	US 2000-710349	20001113
PRIORITY APPLN. INFO.:				
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			EP 1991-310632	A3 19911119
			JP 1991-354242	A3 19911120
			US 1993-44561	B3 19930407
			US 1995-401827	A1 19950310
			US 1995-402296	A1 19950310

OTHER SOURCE(S): MARPAT 118:225657

AB Cyclosporin immunoassay and reagents are disclosed. Interfering cross-reactive material (e.g. cyclosporin metabolites) is inactivated by preparing and using monoclonal antibody to the interfering material. Cyclosporin is conjugated to an immunogenic carrier or label through a linking group at an Ala N of the cyclic backbone of the cyclosporin for preparing antibodies and label reagents for the assay. Atiocyclosporin is conjugated to an immunogenic carrier to prepare antibodies capable of recognizing interfering cross-reactive material but not cyclosporin or cyclosporin conjugates. An EMIT (enzyme multiplied immunoassay techniques) assay was performed on whole blood samples using monoclonal antibody prepared with cyclosporin A bound, through linking group -CH₂(p-Ph)CONHCH₂CONHCH₂CO-, at the Ala N of cyclosporin A amino acid residue number 7 and 8 to keyhole limpet hemocyanin (KLH) as well as glucose-6-phosphate dehydrogenase conjugated with activated cyclosporin A bound, through linking group -(CH₂)₂OCONHCH₂CO-, at the Ala N of cyclosporin A residue 7. Monoclonal antibodies were also prepared against cyclosporin A metabolite M1 using atiocyclosporin conjugated to KLH through an oxyacetic acid imino-extended linking group. The anti-M1 antibody was used in the EMIT assay to inactivate interfering metabolite. Preparation of the above conjugates as well as many others are described.

IT 143676-50-0P

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 (Preparation); RACT (Reactant or reagent)

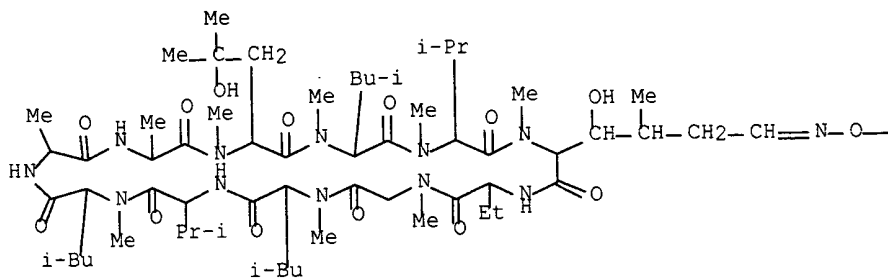
(preparation and reaction of, for preparing monoclonal antibodies to cyclosporin A metabolite M1)

RN 143676-50-0 HCAPLUS

CN Cyclosporin A, 3-(4-hydroxy-N-methyl-L-leucine)-6-[(3R,4R)-6-[(carboxymethoxy)imino]-3-hydroxy-N,4-dimethyl-1-norleucine]- (9CI) (CA

INDEX NAME)

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—CH₂—CO₂H

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143676-64-6P 143676-65-7P

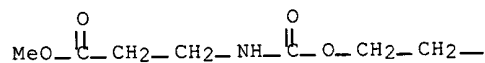
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(preparation and reaction of, in cyclosporin A-glucose-6-phosphate
dehydrogenase conjugate preparation for cyclosporin immunoassay reagent)

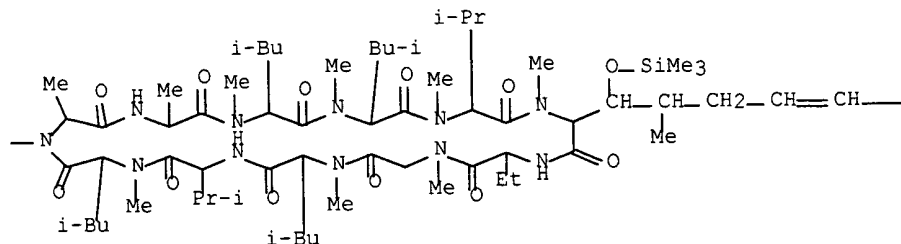
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CN Cyclosporin A, N-(2-hydroxyethyl)-6-[(3R,4R,6E)-6,7-didehydro-N,4-dimethyl-
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N-carboxy-β-alanine methyl ester (9CI) (CA INDEX NAME)

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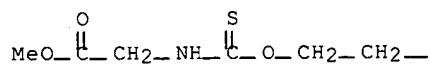


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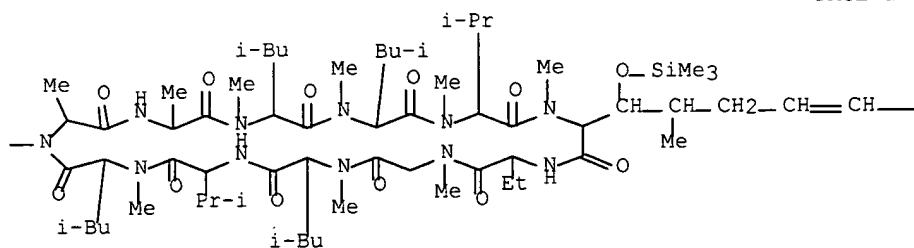
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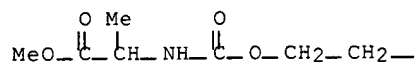


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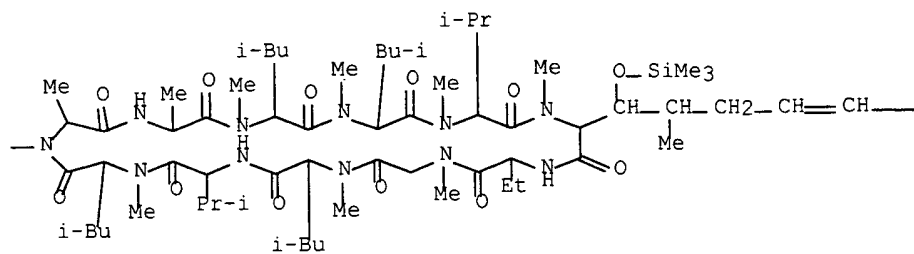
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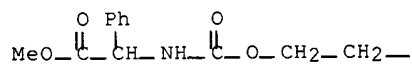


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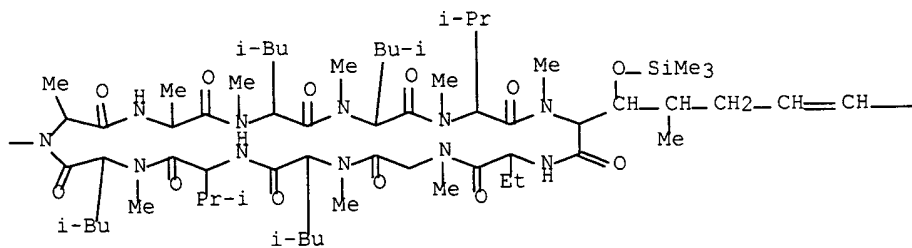
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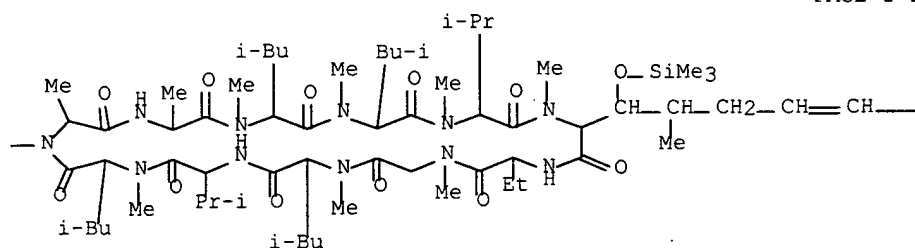
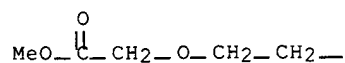


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— Me

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— Me

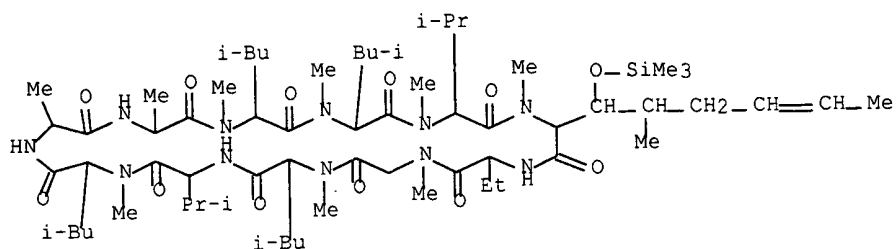
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RL: RCT (Reactant); SPN (Synthetic preparation); PREP
 (Preparation); RACT (Reactant or reagent)

(preparation and reaction of, in cyclosporin A-hemocyanin conjugates preparation for immunogen for cyclosporin immunoassay)

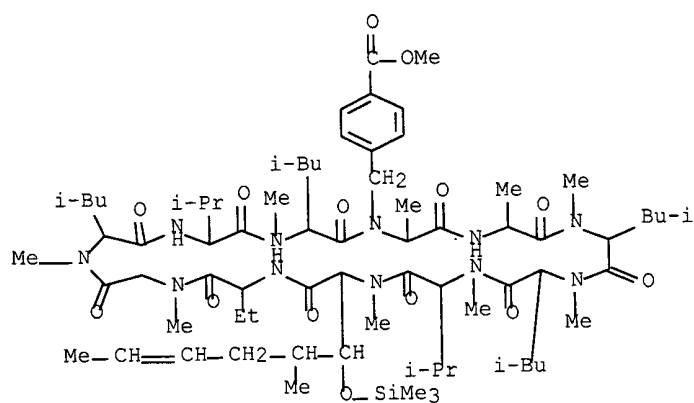
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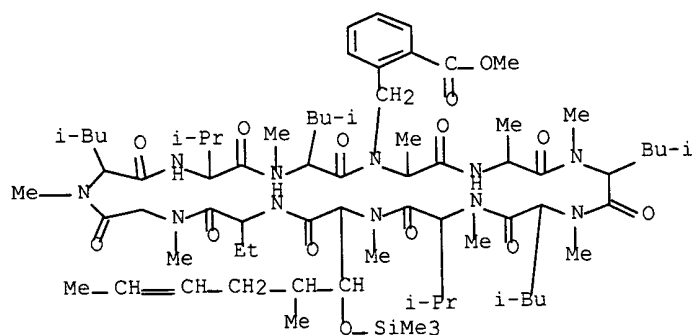
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CN Cyclosporin A, N-[[4-(methoxycarbonyl)phenyl]methyl]-6-[(3R,4R,6E)-6,7-didehydro-N,4-dimethyl-3-[(trimethylsilyl)oxy]-L-2-aminooctanoic acid]-(9CI) (CA INDEX NAME)



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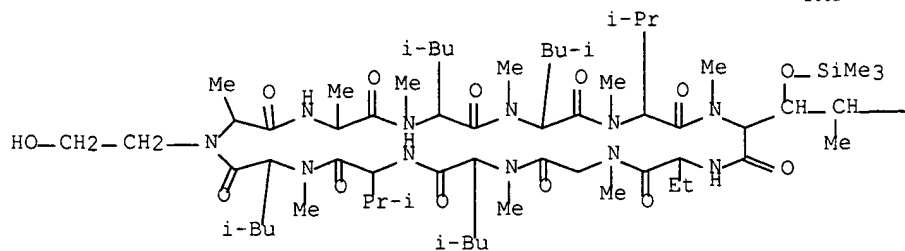
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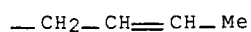
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CN Cyclosporin A, N-(2-hydroxyethyl)-6-[(3R,4R,6E)-6,7-didehydro-N,4-dimethyl-3-[(trimethylsilyl)oxy]-L-2-aminooctanoic acid]-(9CI) (CA INDEX NAME)

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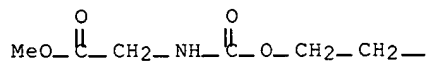


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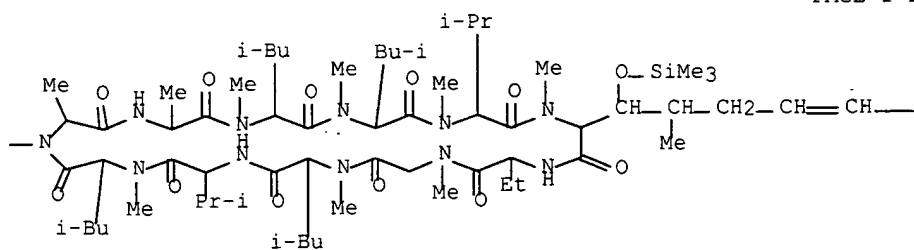


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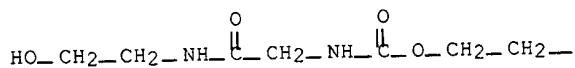


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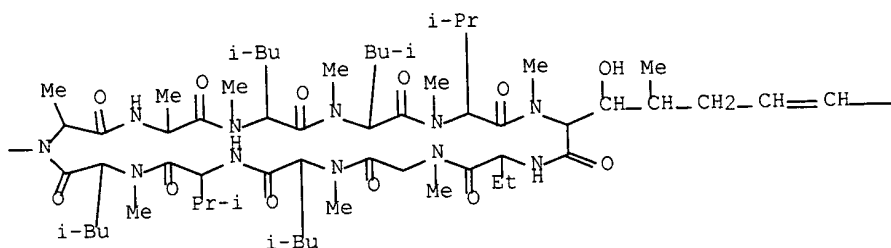


RN 143676-58-8 HCAPLUS
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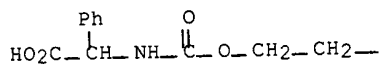
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143676-48-6DP, glucose-6-phosphate dehydrogenase conjugates
143676-49-7DP, glucose-6-phosphate dehydrogenase conjugates
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RL: SPN (Synthetic preparation); **PREP (Preparation)**
 (preparation of, for cyclosporin immunoassay)

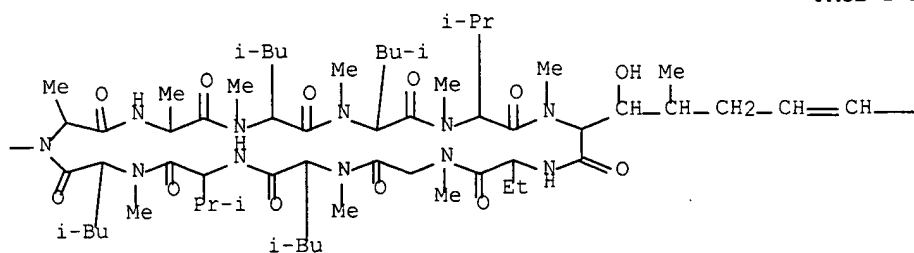
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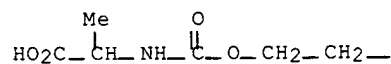


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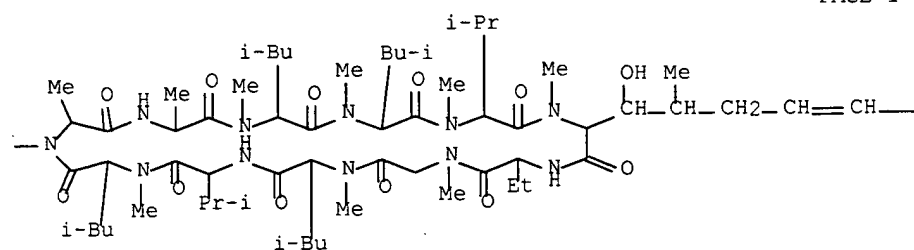
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 N-carboxy-L-alanine (9CI) (CA INDEX NAME)

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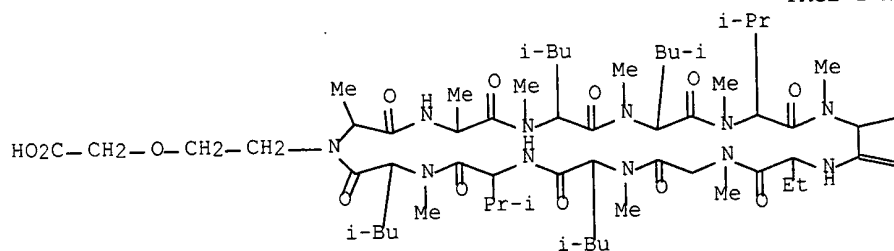


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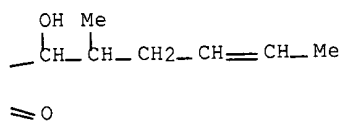
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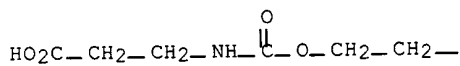


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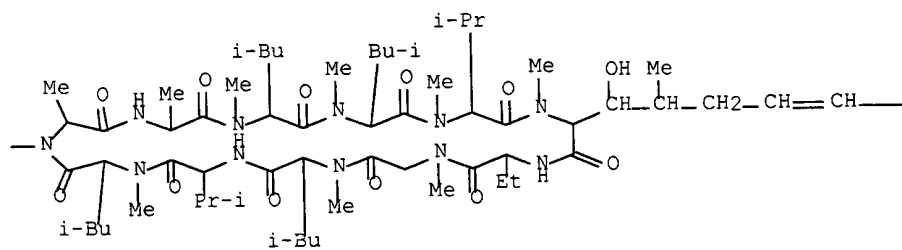


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 N-carboxy-β-alanine (9CI) (CA INDEX NAME)

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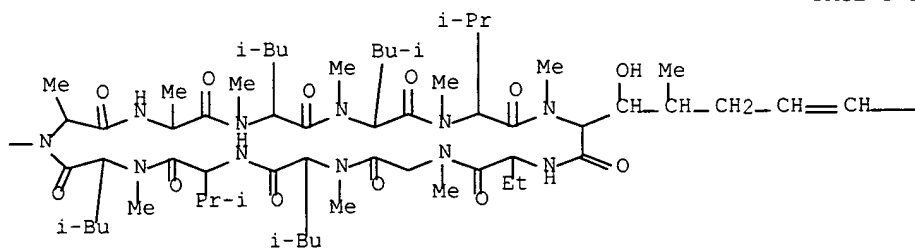
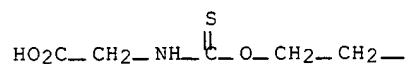


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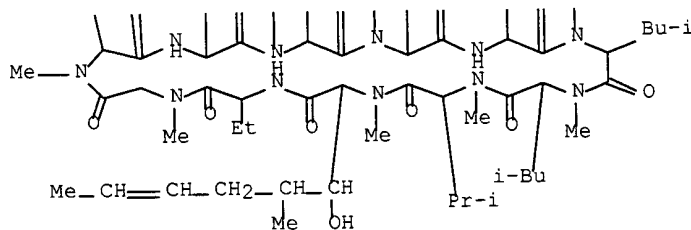
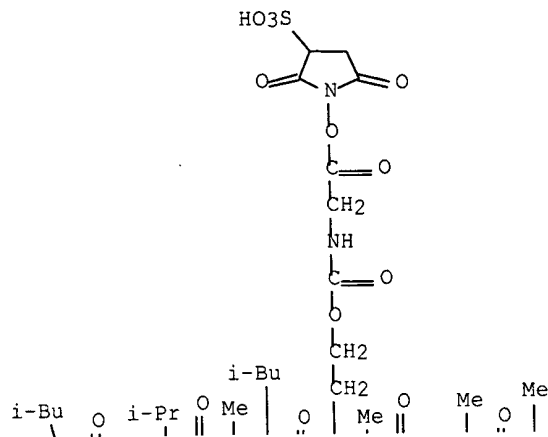
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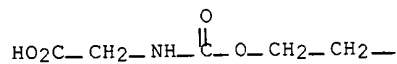
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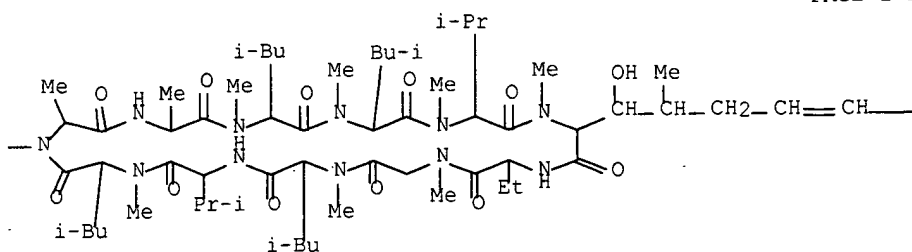
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IT 143676-40-8DP, keyhole limpet hemocyanin conjugates
143676-53-3DP, keyhole limpet hemocyanin conjugates
143676-55-5DP, keyhole limpet hemocyanin conjugates
143676-59-9DP, keyhole limpet hemocyanin conjugates
143676-66-8DP, keyhole limpet hemocyanin conjugates
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of, for immunogen for cyclosporin immunoassay)
 RN 143676-40-8 HCAPLUS
 CN Cyclosporin A, N-(2-hydroxyethyl)-, (1→N)-ester with
 N-carboxyglycine (9CI) (CA INDEX NAME)

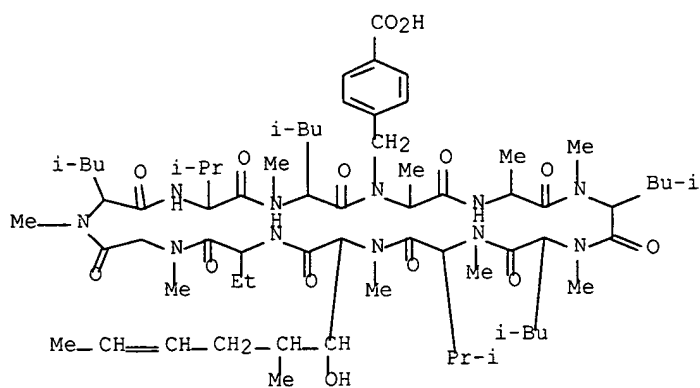




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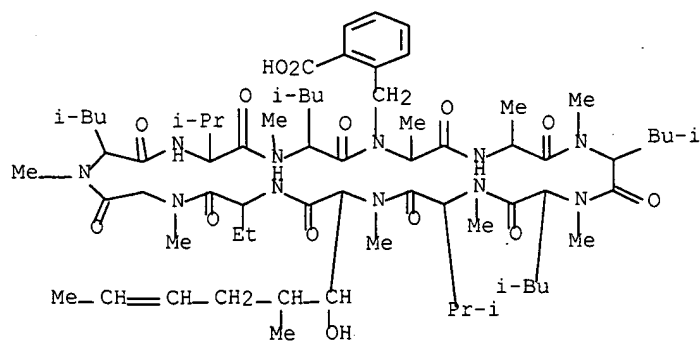
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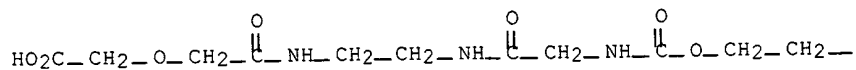
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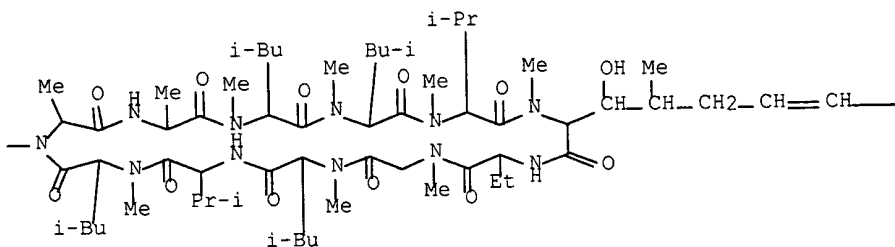


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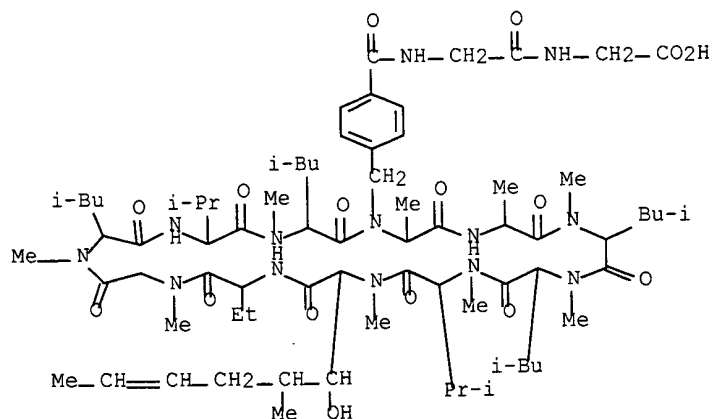
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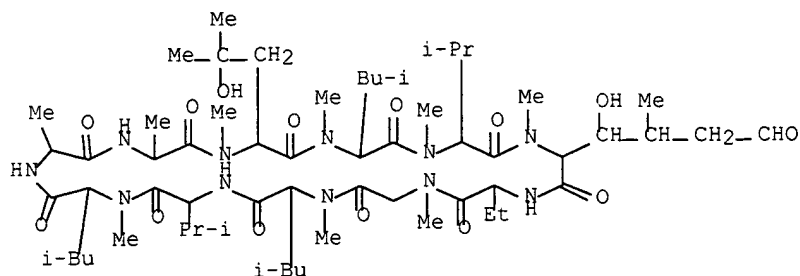
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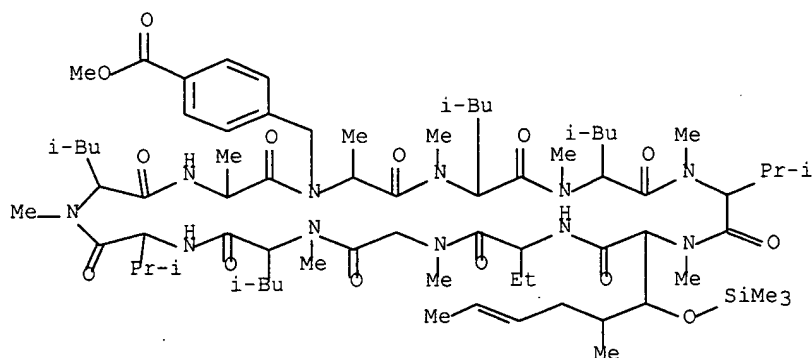
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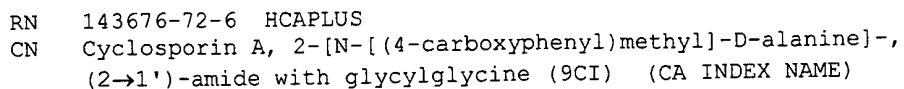
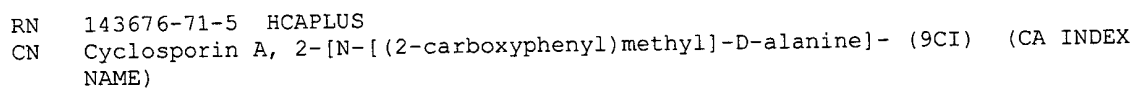
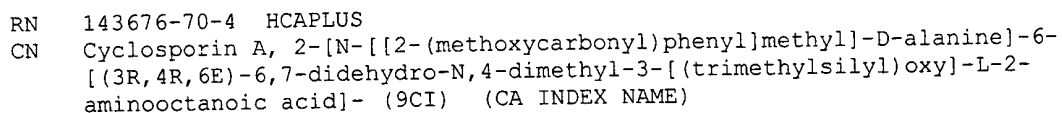
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143676-70-4P 143676-71-5P 143676-72-6P
RL: SPN (Synthetic preparation); PREP (Preparation)
(preparation of, for preparing monoclonal antibodies to cyclosporin A
metabolite M1)
RN 143676-67-9 HCAPLUS
CN Cyclosporin A, 3-(4-hydroxy-N-methyl-L-leucine)-6-[(3R,4R)-3-hydroxy-N,4-
dimethyl-6-oxo-L-norleucine]- (9CI) (CA INDEX NAME)



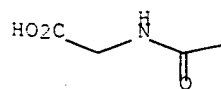
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[(3R,4R,6E)-6,7-didehydro-N,4-dimethyl-3-[(trimethylsilyl)oxy]-L-2-
aminooctanoic acid]- (9CI) (CA INDEX NAME)



RN 143676-69-1 HCAPLUS
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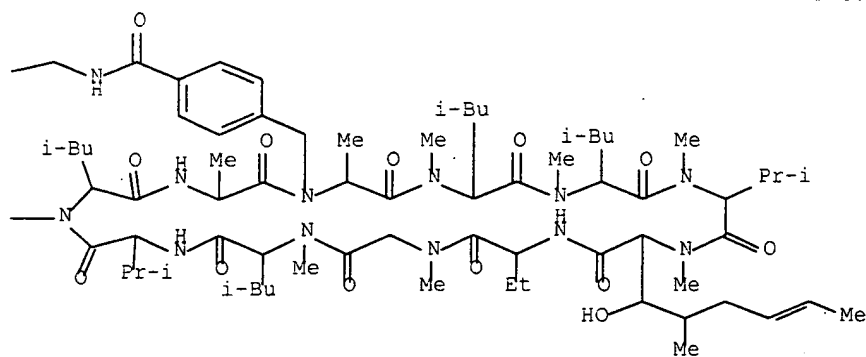


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